I hereby certify that this correspondence is being deposited with the U.S. Postal Service as Express Mail, Airbill No. EV 323 765 795 US, in an envelope addressed to: MS Patent Application, Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450, on the date/shown below.

Signature:

Dated: March 16, 2004

(Jran Quintero)

Docket No.: 29915/00281C

(PATENT)

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Patent Application of: Yan et al.

Application No.: Not Yet Assigned

Art Unit: Not Yet Assigned

Filed: March 16, 2004 (herewith)

Examiner: Not Yet Assigned

For: SUBSTRATES AND ASSAYS FOR BETA-

SECRETASE ACTIVITY

This application is filed as a Divisional application of U.S. Patent Application No. 09/908,943, which was filed July 19, 2001

INFORMATION DISCLOSURE STATEMENT

MS Patent Application Commissioner for Patents P.O. Box 1450

Alexandria, VA 22313-1450

Dear Sir:

Attached herewith are Form 1449A/PTO forms that were submitted in parent application U.S. Patent Application No. 09/908,943. These copies are hereby submitted by the Applicants for consideration in connection with the above-identified patent application in compliance with 37 C.F.R. §1.97 and the duty of disclosure under 37 C.F.R. §1.56.

In accordance with 37 C.F.R. §1.98(d)(1), copies of the references cited on the Form 1449A/PTO are not attached hereto because those references have previously been provided in U.S. Patent Application No. 09/908,943, which was filed July 19, 2001. The present application is a divisional of the aforementioned application, which is relied on for an earlier effective filing date under 35 U.S.C. §120. However, should the Examiner require copies of one or more of the references cited in the prior application, Applicants will furnish the documents upon request.

Application No.: Not Yet Assigned Information Disclosure Statement to Accompany New Divisional Application

of U.S. Serial No. 09/908,943

This information disclosure statement is not intended to be an admission that a search has been made, that other relevant art does not exist, or that any of the information disclosed herein constitutes prior art under 35 U.S.C. §102 or 35 U.S.C. §103.

This statement and the Form 1449A/PTO are being submitted with the application filing papers and as such are submitted before receipt of a first Office Action in the above-identified patent application. Accordingly, it is submitted that no fee is due in this matter under 37 C.F.R. §1.97(b). However, if it is determined that any appropriate fee is due, please charge Deposit Account Number 13-2855. A duplicate of this paper is enclosed.

Dated: March 16, 2004

Respectfully submitted,

Nabeela R. McMillian, Ph.D. Registration No.: 43,363

MARSHALL, GERSTEIN & BORUN LLP

Docket No.: 29915/00281C

233 S. Wacker Drive, Suite 6300

Sears Tower

Chicago, Illinois 60606-6357

(312) 474-6300

Attorney for Applicant

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

| Applicants: Yan et al. |) I hereby certify that this paper is |
|--------------------------------|---|
| Serial No.: 09/908,943 | being deposited with the UnitedStates Postal Service as first class |
| Filed: July 19, 2001 |) mail, postage prepaid, in an envelop) addressed to: Commissioner for |
| For: SUBSTRATES AND ASSAYS FOR |) Patents, Washington, DC 20231, or) this date: |
| BETA-SECRETASE ACTIVITY | \mathbf{j} |
| |) November 19, 2001 |
| Group Art Unit: 1645 | 3 10/10/1 |
| Examiner: To be assigned |) Mille Hale |
| |) Nabeela R. McMillian |
| |) Registration No. 43,363 |
| · |) Agent for Applicants |

INFORMATION DISCLOSURE STATEMENT

Commissioner for Patents Washington, D.C. 20231

Sir:

The patents and/or publications listed on the enclosed Form PTO-1449 are submitted pursuant to 37 CFR §§ 1.56, 1.97, and 1.98. Copies of the patents or publications are enclosed as necessary.

This Information Disclosure Statement is not intended to be an admission that a search has been made, that other relevant art does not exist, or that any of the information disclosed herein constitutes prior art under 35 U.S.C. §102 or §103.

This Information Disclosure Statement is submitted before receipt of a first Office action on the merits, and consequently should be considered by the Patent Office without payment of a fee. See 37 C.F.R. §1.97(b). However, please charge any necessary fees due in connection with this Information Disclosure Statement to Deposit Account No. 13-2855. A copy of this paper is enclosed herewith.

Respectfully submitted,

MARSHALL, GERSTEIN & BORUN 6300 Sears Tower 233 South Wacker Drive Chicago, Illinois 60606-6402 (312) 474-6300

By

Nabeela R. McMillian

Reg. No: 43,363 Agent for Applicants

November 19, 2001

Form PTO-1449 (Modified) U.S. Department of Commerce Atty. Docket No. Serial No. Patent and Trademark Office 29915/00281A 09/908,843 Applicant Yan et al.

INFORMATION DISCLOSURE STATEMENT

(Use several sheets if necessary)

Filing Date Group 1645 July 19, 2001

| | | U.S. PAT | TENT DOCUMENTS | | | |
|-----------------------|--------------------|---------------|----------------|-------|----------|----------------------------|
| *Examiner Initials | Document Number | Issue Date | Name | Class | Subclass | Filing Date If Appropriate |
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| | | F | OREIGN PAT | TENT DOCUM | MENTS | | | |
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| | B1 | WO 00/17369 | 30/03/2000 | WIPO | | | | |

| | OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, etc.) |
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| C1 | Haass et al., "Processing of β-Amyloid Precursor Protein in Microglia and Astrocytes Favors an Internal Localization over Constitutive Secretion", The Journal of Neuroscience, 11(12):3783-3793 |
| C2 | Kang et al., "The precursor of Alzheimer's disease amyloid A4 protein resembles a cell-surface receptor", Nature, 325:733-736 (1987) |
| C3 | Kitaguchi et al., "Novel precursor of Alzheimer's disease amyloid protein shows protease inhibitory activity", Nature, 331:530-532 (1988) |
| C4 | Lin et al., "Inhibition of Cathepsin D by Synthetic Oligonucleotides", The Journal of Biological Chemistry, 254:11875-11883 (1979) |
| C5 | Lin et al., "Human aspartic protease memapsin 2 cleaves the β -secretase site of β -amyloid precursor protein", PNAS, 97:1456-1460 (2000) |
| C6 | Ponte et al., "A new A4 amyloid mRNA contains a domain homologous to serine proteinase inhibitors", Nature, 331:525-527 (1988) |

| EXAMINER | | DATE CONSIDERED |
|----------|---|-----------------|
| | • | DATE CONSIDERED |
| | | II. |

^{*}EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

| Form PT |)-1449 (Modified) | |
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U.S. Department of Commerce Patent and Trademark Office Atty. Docket No. 29915/00281A

Serial No. 09/908,843

Applicant

Yan et al.

Filing Date
July 19, 2001

Group 1645

INFORMATION DISCLOSURE STATEMENT

(Use several sheets if necessary)

| | | OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, etc.) |
|---|-----|--|
| | C7 | Schechter et al., "On the Active Site of Proteases, III. Mapping the Active Site of Papain; Specific Peptide Inhibitors of Papain", Biochemical and Biophysical Research Communications, 32:898-902 (1968) |
| , | C8 | Schechter et al., "On the Size of the Active Site in Proteases, I. Papain", Biochemical and Biophysical Research Communications, 27:157-162 (1967) |
| | C9 | Sinha et al., "Cellular mechanisms of β-amyloid production and secretion", Proc. Nat'l. Acad. Sci. (USA), 96:11049-11053 (1999) |
| | C10 | Tanzi et al., "Protease inhibitor domain encoded by an amyloid protein precursor mRNA associated with Alzheimer's disease", Nature, 331:528-530 (1988) |
| | C11 | Vassar et al., "β-Secretase Cleavage of Alzheimer's Amyloid Precursor Protein by the Transmembrane Aspartic Protease BACE", Science, 286:735-741 (1999) |
| | C12 | Yan et al., "Membrane-anchored aspartyl protease with Alzheimer's disease β - secretase activity", Nature, 402:533-537 (1999) |
| | C13 | Younkin, "Processing of the Alzheimer's Disease β A4 Amyloid Protein Precursor (APP)", Brain Pathology, 1:253-262 (1991) |

EXAMINER

DATE CONSIDERED

*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609; Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

I hereby certify that this correspondence is being deposited with the U.S. Postal Service with sufficient postage as First Class Mail, in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450, on the date shown below.

Dated: May 6, 2003

Signature: Nahada P. Mahadilian

Docket No.: 29915/00281A

(PATENT)

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Patent Application of: Yan et al.

Application No.: 09/908,943

Group Art Unit: 1652

Filed: July 19, 2001

Examiner: Charles L. Patterson, Jr.

For:

SUBSTRATES AND ASSAYS FOR

β-SECRETASE ACTIVITY

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT (IDS)

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

Dear Sir:

Pursuant to 37 C.F.R. 1.56, the attention of the Patent and Trademark Office is hereby directed to the references listed on the attached Form PTO - 1449. It is respectfully requested that the information be expressly considered during the prosecution of this application, and that the references be made of record therein and appear among the "References Cited" on any patent to issue therefrom.

All patents and publications in the attached Form PTO - 1449 are supplied herewith. It is not intended to constitute an admission that any patent, publication or other information referred to therein is "prior art" for the instant application under any section of 35 U.S.C. §102 or 35 U.S.C. §103.

In accordance with 37 C.F.R. 1.97(g), the filing of this Information Disclosure Statement shall not be construed to mean that a search has been made or that no other material information as defined in 37 C.F.R. 1.56(a) exists. It is submitted that the Information Disclosure Statement is in compliance with 37 C.F.R. 1.98 and the Examiner is respectfully requested to consider the listed references.

Application No.: 09/908,943 Docket No.: 29915/00281A

In accordance with 37 C.F.R. §1.97(e)(2), individuals designated in § 1.56(c) were not aware of the reference cited herewith more than three months prior to the filing of this Information Disclosure Statement. Therefore, no fees are believed to be necessary, however, the Commissioner is hereby authorized to charge any deficiency in the fees filed or credit any overpayments to our Deposit Account No. 13-2855. A duplicate copy of this paper is enclosed.

Dated: May 6, 2003

Respectfully submitted,

Nabeela R. McMillian

Registration No.: 43,363 MARSHALL, GERSTEIN & BORUN

233 S. Wacker Drive, Suite 6300

Sears Tower

Chicago, Illinois 60606-6357

(312) 474-6300

Attorneys for Applicants

| Form PTO-1449 (Modified) U.S. Depart Patent and | ment of Commerce 1 Trademark Office 29915/00281A Serial No 09/908,943 | |
|---|---|-----|
| SUPPLEMENTAL INFORMATION DISCL | Applicant . | · · |
| STATEMENT | July 19, 2001 Group 1652 | |

| U.S. PATENT DOCUMENTS | | | | | | |
|-----------------------|--------------------|------------|------|-------|----------|----------------------------|
| *Examiner Initials | Document Number | Issue Date | Name | Class | Subclass | Filing Date if Appropriate |
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|-----------------------|-----------------|---------------------|-----------|-------|----------|--------------------|----|
| *Examiner Initials | Document Number | Publication Date | Country | Class | Subclass | Translation Yes | No |
| | WO 02/094985 | 11/28/2002 | PCT | | | · | |

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| | OTHER DOCUMENTS (In | cluding Author, Tit | le, Date, Pertinent Pages, etc.) | |
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Signature

Dated: May 28, 2003

Nabeela R-McMillian

Docket No.: 29915/00281A.US

(PATENT)

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Patent Application of: Yan et al.

Application No.: 09/908,943 Group Art Unit: 1652

Filed: July 19, 2001 Examiner: Charles L. Patterson, Jr.

For: SUBSTRATES AND ASSAYS FOR

β-SECRETASE ACTIVITY

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT (IDS)

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

Dear Sir:

Pursuant to 37 CFR 1.56, the attention of the Patent and Trademark Office is hereby directed to the references listed on the attached Form PTO - 1449. It is respectfully requested that the information be expressly considered during the prosecution of this application, and that the references be made of record therein and appear among the "References Cited" on any patent to issue therefrom.

All patents and publications in the attached Form PTO - 1449 are supplied herewith. It is not intended to constitute an admission that any patent, publication or other information referred to therein is "prior art" for the instant application under any section of 35 U.S.C. §102 or 35 U.S.C. §103.

In accordance with 37 CFR 1.97(g), the filing of this Information Disclosure Statement shall not be construed to mean that a search has been made or that no other material information as defined in 37 CFR 1.56(a) exists. It is submitted that the Information Disclosure Statement is in compliance with 37 CFR 1.98 and the Examiner is respectfully requested to consider the listed references.

Application No.: 09/908,943 Docket No.: 29915/00281A.US

This Information Disclosure Statement is being filed after the receipt of a first office action. Therefore, a fee pursuant to 37 C.F.R. §1.17(p) for the filing of the Information Disclosure Statement is attached herewith. No additional fees are believed to be necessary, however, the Commissioner is hereby authorized to charge any deficiency in the fees filed or credit any overpayments to our Deposit Account No. 13-2855. A duplicate copy of this paper is enclosed.

Dated: May 28, 2003

Respectfully submitted,

Nabeela R. McMillian

Registration No.: 43,363

MARSHALL, GERSTEIN & BORUN

233 S. Wacker Drive, Suite 6300

Sears Tower

Chicago, Illinois 60606-6357

(312) 474-6300

Agent for Applicants

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|--------------------------|---|--|--------------------------|-------------|
| Form PTO-1449 (Modified) | U.S. Department of Commerce Patent and Trademark Office | Attorney Docket No. 29915/00281A.US | Serial No. 09/908,943 | |
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| SUPPLEMENTAL INFORM | | 2 | Group | |
| STATEM | IENT | Filing Date Tuly 10 2001 | 1652 | |

| U.S. PATENT DOCUMENTS | | | | | | | | | |
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| *Examiner Initials | | Document Number | Issue Date | Name | Class | Subclass | Filing Date if Appropriate | | |
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| | FOREIGN PATENT DOCUMENTS | | | | | | | |
|-----------------------|--------------------------|-----------------|---------------------|---------|-------|----------|--------------------|----|
| *Examiner Initials | | Document Number | Publication Date | Country | Class | Subclass | Translation Yes | No |
| | | WO 92/07068 | 04/30/1992 | PCT | | | | |

| OTHER DOCU | MENTS (Including Au | thor, Title, Date, l | Pertinent Pages | , etc.) | |
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EXAMINER

DATE CONSIDERED

I hereby certify that this correspondence is being deposited with the U.S. Postal Service with sufficient postage as First Class Mail, in an envelope addressed to: Commissioner for Patents, P.O. Box 1450, Alexandria, VA 22313-1450, on the date shown below.

Signature

Dated: February 3, 2004

Nabeela R. McMillian

Docket No.: 29915/00281A.US

(PATENT)

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

In re Patent Application of: Yan et al.

Application No.: 09/908,943 Group Art Unit: 1652

Filed: July 19, 2001 Examiner: Charles L. Patterson, Jr.

For: SUBSTRATES AND ASSAYS FOR BETA-

SECRETASE ACTIVITY

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT (IDS)

Commissioner for Patents P.O. Box 1450 Alexandria, VA 22313-1450

Dear Sir:

Pursuant to 37 CFR 1.56, the attention of the Patent and Trademark Office is hereby directed to the references listed on the attached Form PTO - 1449. It is respectfully requested that the information be expressly considered during the prosecution of this application, and that the references be made of record therein and appear among the "References Cited" on any patent to issue therefrom.

All patents and publications in the attached Form PTO - 1449 are supplied herewith. It is not intended to constitute an admission that any patent, publication or other information referred to therein is "prior art" for the instant application under any section of 35 U.S.C. §102 or 35 U.S.C. §103.

In accordance with 37 CFR 1.97(g), the filing of this Information Disclosure Statement shall not be construed to mean that a search has been made or that no other material information as defined in 37 CFR 1.56(a) exists. It is submitted that the Information Disclosure Statement is in compliance with 37 CFR 1.98 and the Examiner is respectfully requested to consider the listed references.

Application No.: 09/908,943 Docket No.: 29915/00281A.US

This Information Disclosure Statement is being filed after the receipt of a first office action. It is filed to make of record publications discussed in an International Search Report in a related International patent application. A fee pursuant to 37 C.F.R. §1.17(p) for the filing of the Information Disclosure Statement is attached herewith. No additional fees are believed to be necessary, however, the Commissioner is hereby authorized to charge any deficiency in the fees filed or credit any overpayments to our Deposit Account No. 13-2855. A duplicate copy of this paper is enclosed.

Dated: February 3, 2004

Respectfully submitted,

By Mullian Nabeela R. McMillian

Registration No.: 43,363

MARSHALL, GERSTEIN & BORUN LLP

233 S. Wacker Drive, Suite 6300

Sears Tower

Chicago, Illinois 60606-6357

(312) 474-6300

Attorneys for Applicants

| · · · · · · · · · · · · · · · · · · · | | Sheet 1 of 1 |
|---|---------------------|--------------|
| Form PTO-1449 (Modified) | Attorney Docket No. | Serial No |
| U.S. Department of Commerce Patent and Trademark Office | 29915/00281A.US | 09/908,943 |
| | Applicant | . • |
| SUPPLEMENTAL INFORMATION DISCLOSURE | Yan et al. | • |
| STATEMENT | Filing Date | Group |
| | July 19, 2001 | 1652 |

| | U.S. PATENT DOCUMENTS | | | | | | | |
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| *Examiner Initials | | Document Number | Issue Date | Name | Class | Subclass | Filing Date if Appropriate | |
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| *Examiner Initials | Document Number | Publication Date | Country | Class | Subclass | Translation Yes | No |
| · | EP 0 855 444 | 07/29/1998 | EPO | | | | |
| | WO 00/47618 | 08/17/2000 | PCT | | | | |
| · | WO 00/58479 | 10/05/2000 | PCT | | 6 | | |

| | | OTHER DOCUMENTS (Including Author, Title, Date, Pertinent Pages, etc.) | * | : |
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| | T . | Bailey et al., Pro. Science, 3:2129-2143 (1994) | | ····· |
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| | | Sinha et al., Nature, 402:537-540 (1999) | | • |
| • . | | Yang et al., Acta Cryst., D55:625-630 (1999) | | • • • • |